

CLAIMS

What is claimed is:

1. A polynucleotide capable of inducing methylation at a target nucleotide sequence comprising:
 - 5 an oligonucleotide imprinting element, wherein said imprinting element is double stranded and has a first strand and a second strand complementary to said first strand, said first strand comprising at least one m5CG sequence paired with an unmethylated CG sequence on said second strand or at least one m5CN1G sequence paired with an unmethylated CN2G sequence on said second strand, wherein N1 is any nucleotide, and
10 N2 is a nucleotide that pairs with N1; and
 - a single stranded oligonucleotide guiding element comprising at least one m5CG or at least one m5CN3G sequence, wherein N3 is any nucleotide and said guiding element is complementary to a target nucleotide sequence; and
 - 15 wherein said imprinting element and said guiding element are operably linked to form a polynucleotide capable of inducing methylation at said target nucleotide sequence.
2. The polynucleotide of claim 1 wherein said first strand and said second strand of said imprinting element are linked through a covalent linkage.
3. The polynucleotide of claim 2 wherein said first strand and said second strand of said imprinting element are linked by one or more nucleotide residues.
- 20 4. The polynucleotide of claim 3 wherein said first strand and said second strand of said imprinting element are linked by a single thymidine residue.
5. The polynucleotide of claim 1 wherein said first strand of said imprinting element comprises at least two m5CG sequences, two m5CN1G sequences, or one m5CG sequence and one m5CN1G sequence.

6. The polynucleotide of claim 5 wherein said first strand of said imprinting element comprises the sequence: 5'-m5CGpN1m5CG-3'.
7. The polynucleotide of claim 6 wherein said first strand of said imprinting element comprises the sequence: 5'-m5CGpTpm5CG-3'.
- 5 8. The polynucleotide of claim 7 wherein said imprinting element comprises the sequence: 5'-CGpApCG-T-m5CGpTpm5CG-3' (SEQ ID NO:zz).
9. The polynucleotide of claim 1 wherein said guiding element is about 15 to about 30 nucleotides in length.
10. The polynucleotide of claim 9 wherein said guiding element is a 22-nucleotide oligomer having the sequence 5'-AGCC^mCGGG^mCTGGGAGGAGT^mCGG-3' (SEQ ID NO:zz).
11. The polynucleotide of claim 10 having the sequence 5'-CGACGT^mCGpTp^mCGAGCC^mCGGG^mCTGGGAGGAGT^mCGG-3' (SEQ ID NO:zz).
12. The polynucleotide of claim 1 wherein said guiding element is operably linked to the 3' end of said imprinting element.
- 15 13. The polynucleotide of claim 1 wherein said target nucleotide sequence is in a gene.
14. The polynucleotide of claim 13 wherein said target nucleotide sequence is in a gene regulatory region.
15. The polynucleotide of claim 14 wherein said gene encodes a protein of unknown function.
- 20 16. The polynucleotide of claim 15 wherein the gene is a disease gene.
17. The polynucleotide of claim 16 wherein the disease is cancer.
18. The polynucleotide of claim 14 wherein said target nucleotide sequence is in a gene regulatory region of human Igf2.

19. A composition comprising the polynucleotide of claim 1 and one or more additional components.
20. The composition of claim 19 wherein said additional component is one that facilitates entry of the polynucleotide into a cell.
- 5 21. The composition of claim 20 wherein said additional component comprises one or more lipids.
22. The composition of claim 21 wherein said one or more lipids comprise the cationic lipids N-[1-(2,3-dioleoyloxy)propyl]-n,n,n-trimethylammonium chloride (DOTMA), dioleoyl phosphatidylethanolamine (DOPE) and/or dioleoyl phosphatidylcholine (DOPC).
- 10 23. The composition of claim 19 wherein said additional component is a physiologically acceptable carrier.
24. A method for inducing methylation at a target nucleotide sequence in a cell comprising introducing the polynucleotide of claim 1 into a cell comprising said target nucleotide sequence, thereby inducing methylation at said target nucleotide sequence.
- 15 25. The method of claim 24 wherein said cell is a mammalian cell, a plant cell, or a prokaryotic cell.
26. The method of claim 25 wherein said cell is a human cell.
27. The method of claim 24 wherein said polynucleotide is introduced into said cell *in vivo*.
28. The method of claim 24 wherein said polynucleotide is introduced into said cell *ex vivo*.
- 20 29. The method of claim 24 wherein said polynucleotide is introduced as a composition comprising a lipid.
30. The method of claim 24 additionally comprising determining a phenotypic change associated with methylation at said target nucleotide sequence after introduction of said polynucleotide.

31. The method of claim 30 wherein said target nucleotide sequence is in a gene encoding a protein of unknown function.

32. The method of claim 24 additionally comprising producing an organism from said cell, wherein said target sequence is in a gene, and the organism either does not express said gene or expresses said gene at a reduced level compared to a normal organism.

33. The method of claim 24 wherein said target nucleotide sequence is in a disease gene.

34. The method of claim 33 wherein said disease is cancer.